

Amendments to the Claims:

This listing of claims will replace all prior versions and listings of claims in the application.

Listing of Claims:

1. (Currently Amended) A method of making an array-comprising reversibly immobilized building blocks, the method comprising:
applying building block molecules to a solid support in a plurality of regions-spots, the spots each region:
comprising 2, 3, 4, or 5, or 6 different building block molecules;
being a contiguous portion of the surface of the solid support with the different building block molecules distributed randomly and evenly throughout the contiguous region; and
having the shape of a spot;
independently reversibly immobilizing the different block molecules on the solid support in the regions-spots;
producing an array comprising a candidate artificial receptor, a lead artificial receptor, a working artificial receptor, or a combination thereof;
wherein:
a first region spot comprises a first combination of block molecules and a second region spot comprises a second combination of block molecules;
at least one of the block molecules is naïve with respect to a test ligand; and
the spots-of-immobilized-blocks-on-the-solid-support-form-a-heterogeneous building-block-array
each building block molecule comprises a framework and n recognition elements and is independently of the formula:
$$\text{framework-(recognition element)}_n$$

in which:
n=1, 2, or 3; each recognition element is independently covalently coupled to the framework; and the framework comprises a functional group effective for covalent coupling to a

linker, a structural moiety or functional group effective for reversible coupling to a support, or a combination thereof;

the framework is alkyl, substituted alkyl, cycloalkyl, heterocyclic, substituted heterocyclic, aryl alkyl, aryl, heteroaryl, or heteroaryl alkyl; substituted with 1 to 4 functional groups;

the functional groups independently being carboxyl, amine, hydroxyl, phenol, carbonyl, or thiol;

each recognition element is independently a 1-12 carbon alkyl, substituted alkyl, cycloalkyl, heterocyclic, substituted heterocyclic, aryl alkyl, aryl, heteroaryl, or heteroaryl alkyl moiety; substituted with a group with a property of positive charge, negative charge, acid, base, electron acceptor, electron donor, hydrogen bond donor, hydrogen bond acceptor, free electron pair, π electrons, charge polarization, hydrophilicity, or hydrophobicity.

2. (Currently Amended) The method of claim 1, further comprising:
mixing the different block molecules; and
wherein applying comprises applying employing the mixture of building block molecules to the solid support in at least one of forming the regions plurality of spots.
3. (Previously presented) The method of claim 1, wherein applying comprises piezoelectric spotting, pin spotting, electromagnetic spotting, or photolithography.
4. (Original) The method of claim 1, wherein the solid support comprises a glass plate or microscope slide.
- 5-7. (canceled)
8. (Currently Amended) A method of making an artificial receptor, the method comprising:
applying building block molecules to a region on a solid support, the region:
comprising 2, 3, 4, 5, or 6 different building block molecules; and

being a contiguous portion of the surface of the solid support with the different building block molecules distributed randomly and evenly throughout the contiguous region;
independently reversibly immobilizing the different building block molecules on the support in the region;

wherein:
2 or more of the different building block molecules together forming a candidate artificial receptor, a lead artificial receptor, a working artificial receptor, or a combination thereof;
at least one of the building block molecules is being naïve with respect to a test ligand.
each building block molecule comprises a framework and n recognition elements and is independently of the formula:

framework-(recognition element)_n
in which:
n=1, 2, or 3; each recognition element is independently covalently coupled to the framework; and the framework comprises a functional group effective for covalent coupling to a linker, a structural moiety or functional group effective for reversible coupling to a support, or a combination thereof;

the framework is alkyl, substituted alkyl, cycloalkyl, heterocyclic, substituted heterocyclic, aryl alkyl, aryl, heteroaryl, or heteroaryl alkyl; substituted with 1 to 4 functional groups;

the functional groups independently being carboxyl, amine, hydroxyl, phenol, carbonyl, or thiol;
each recognition element is independently a 1-12 carbon alkyl, substituted alkyl, cycloalkyl, heterocyclic, substituted heterocyclic, aryl alkyl, aryl, heteroaryl, or heteroaryl alkyl moiety; substituted with a group with a property of positive charge, negative charge, acid, base, electron acceptor, electron donor, hydrogen bond donor, hydrogen bond acceptor, free electron pair, π electrons, charge polarization, hydrophilicity, or hydrophobicity.

9. (Currently Amended) The method of claim 8, wherein the region is a region spot.

10-66. (Cancelled).

67. (New) The method of claim 8, further comprising:
mixing the different block molecules; and
wherein applying comprises applying employing the mixture of building block molecules to the solid support in the region.

68. (New) The method of claim 8, wherein applying comprises piezoelectric spotting, pin spotting, electromagnetic spotting, or photolithography.

69. (New) The method of claim 8, wherein the solid support comprises a glass plate or microscope slide.

70. (New) The method of claim 1, wherein each recognition element is independently substituted with a moiety selected from the group consisting of amine, quaternary ammonium, carboxylate, phenol, phosphate, phosphonate, phosphinate, sulphate, sulphonate, thiocarboxylate, hydroxamic acid, sulfoxide, betaine, amine oxide, amide, carboxyl, alcohol, ether, thiol, thioether, ester, thio ester, borane, borate, metal complex, alkyl, alkene, alkyne, aromatic moiety, and plurality thereof.

71. (New) The method of claim 1, wherein a recognition element is substituted with or to form:

protonated phosphate, protonated phosphonate, protonated phosphinate, protonated sulphate, or protonated sulphinate;

alkyl amine, alkyl diamine, heteroalkyl amine, aryl amine, heteroaryl amine, aryl alkyl amine, heterocyclic amine, amidine, hydrazine, urea, trimethyl alkyl quaternary ammonium, dimethyl ethyl alkyl quaternary ammonium, dimethyl alkyl quaternary ammonium, aryl alkyl quaternary ammonium, or pyridinium quaternary ammonium;

alkyl carboxylate, aryl carboxylate, aryl alkyl carboxylate, or thiocarboxylate;
phosphonate or phosphinate;

primary alcohol, secondary alcohol, tertiary alcohol, or aromatic alcohol;
lower alkyl, substituted alkyl, cycloalkyl, aryl alkyl, heteroaryl alkyl, lower alkene, aryl
alkene, unsubstituted aryl, heteroaryl, substituted aryl, aryl alkyl, heteroaryl alkyl, alkyl
substituted aryl, or polyaromatic hydrocarbon; or
a plurality thereof.

72. (New) The method of claim 1, wherein the framework has the formula:



in which:

R₁ is 1-12 carbon alkyl, substituted alkyl, cycloalkyl, heterocyclic, substituted
heterocyclic, aryl alkyl, aryl, heteroaryl, or heteroaryl alkyl;

F₁ and F₂ are independently carboxyl, amine, hydroxyl, phenol, carbonyl, or thiol group;
or are independently 1-12 carbon alkyl, substituted alkyl, cycloalkyl, heterocyclic, substituted
heterocyclic, aryl alkyl, aryl, heteroaryl, heteroaryl alkyl, or inorganic group substituted with
carboxyl, amine, hydroxyl, phenol, carbonyl, or thiol group;

F₃ and F₄ are independently absent, carboxyl, amine, hydroxyl, phenol, carbonyl, or thiol
group; or are independently absent, or 1-12 carbon alkyl, substituted alkyl, cycloalkyl,
heterocyclic, substituted heterocyclic, aryl alkyl, aryl, heteroaryl, heteroaryl alkyl, or inorganic
group substituted with carboxyl, amine, hydroxyl, phenol, carbonyl, or thiol group.

73. (New) The method of claim 72, wherein:

R₁ is 1-6 carbon alkyl, substituted alkyl, cycloalkyl, heterocyclic, substituted
heterocyclic, aryl alkyl, aryl, heteroaryl, or heteroaryl alkyl;

F₁, F₂, F₃, or F₄ are independently 1-6 carbon alkyl, substituted alkyl, cycloalkyl,
heterocyclic, substituted heterocyclic, aryl alkyl, aryl, heteroaryl, heteroaryl alkyl, or inorganic
group substituted with carboxyl, amine, hydroxyl, phenol, carbonyl, or thiol group.

F₃ is absent; or

F₃ and F₄ are absent.

74. (New) The method of claim 1, wherein the framework is:
a natural or synthetic amino acid, an α -hydroxy acid, or a thioic acid; or
a β -amino acids or homo or β analog of a natural amino acid.
75. (New) The method of claim 1, wherein the framework is an amino acid with an amine, hydroxyl, phenol, carboxyl, thiol, thioether, or amidino group on its side chain.
76. (New) The method of claim 1, wherein the framework is a serine, threonine, tyrosine, aspartic acid, glutamic acid, asparagine, glutamine, cysteine, lysine, arginine, or histidine moiety.
77. (New) The method of claim 1, wherein the building block molecule further comprises a linker, and
the linker is a alkyl, substituted alkyl, cycloalkyl, heterocyclic, substituted heterocyclic, aryl alkyl, aryl, heteroaryl, heteroaryl alkyl, ethoxy or propoxy oligomer, or glycoside moiety;
substituted with a carboxyl, alcohol, phenol, thiol, amine, carbonyl, or maleimide group.
78. (Currently Amended) The method of claim 77, wherein the linker is of the formula $(CH_2)_nC(O)-$, with $n=1-16$.
79. (New) The method of claim 8, wherein each recognition element is independently substituted with a moiety selected from the group consisting of amine, quaternary ammonium, carboxylate, phenol, phosphate, phosphonate, phosphinate, sulphate, sulphonate, thiocarboxylate, hydroxamic acid, sulfoxide, betaine, amine oxide, amide, carboxyl, alcohol, ether, thiol, thioether, ester, thio ester, borane, borate, metal complex, alkyl, alkene, alkyne, aromatic moiety, and plurality thereof.

80. (New) The method of claim 8, wherein a recognition element is substituted with or to form:

protonated phosphate, protonated phosphonate, protonated phosphinate, protonated sulphate, or protonated sulphinate;

alkyl amine, alkyl diamine, heteroalkyl amine, aryl amine, heteroaryl amine, aryl alkyl amine, heterocyclic amine, amidine, hydrazine, urea, trimethyl alkyl quaternary ammonium, dimethyl ethyl alkyl quaternary ammonium, dimethyl alkyl quaternary ammonium, aryl alkyl quaternary ammonium, or pyridinium quaternary ammonium;

alkyl carboxylate, aryl carboxylate, aryl alkyl carboxylate, or thiocarboxylate;

phosphonate or phosphinate;

primary alcohol, secondary alcohol, tertiary alcohol, or aromatic alcohol;

lower alkyl, substituted alkyl, cycloalkyl, aryl alkyl, heteroaryl alkyl, lower alkene, aryl alkene, unsubstituted aryl, heteroaryl, substituted aryl, aryl alkyl, heteroaryl alkyl, alkyl substituted aryl, or polyaromatic hydrocarbon; or

a plurality thereof.

81. (New) The method of claim 8, wherein the framework has the formula:



in which:

R₁ is 1-12 carbon alkyl, substituted alkyl, cycloalkyl, heterocyclic, substituted heterocyclic, aryl alkyl, aryl, heteroaryl, or heteroaryl alkyl;

F₁ and F₂ are independently carboxyl, amine, hydroxyl, phenol, carbonyl, or thiol group; or are independently 1-12 carbon alkyl, substituted alkyl, cycloalkyl, heterocyclic, substituted heterocyclic, aryl alkyl, aryl, heteroaryl, heteroaryl alkyl, or inorganic group substituted with carboxyl, amine, hydroxyl, phenol, carbonyl, or thiol group;

F₃ and F₄ are independently absent, carboxyl, amine, hydroxyl, phenol, carbonyl, or thiol group; or are independently absent, or 1-12 carbon alkyl, substituted alkyl, cycloalkyl,

heterocyclic, substituted heterocyclic, aryl alkyl, aryl, heteroaryl, heteroaryl alkyl, or inorganic group substituted with carboxyl, amine, hydroxyl, phenol, carbonyl, or thiol group.

82. (New) The method of claim 81, wherein:

R₁ is 1-6 carbon alkyl, substituted alkyl, cycloalkyl, heterocyclic, substituted heterocyclic, aryl alkyl, aryl, heteroaryl, or heteroaryl alkyl;

F₁, F₂, F₃, or F₄ are independently 1-6 carbon alkyl, substituted alkyl, cycloalkyl, heterocyclic, substituted heterocyclic, aryl alkyl, aryl, heteroaryl, heteroaryl alkyl, or inorganic group substituted with carboxyl, amine, hydroxyl, phenol, carbonyl, or thiol group.

F₃ is absent; or

F₃ and F₄ are absent.

83. (New) The method of claim 8, wherein the framework is:

a natural or synthetic amino acid, an α -hydroxy acid, or a thioic acid; or

a β -amino acids or homo or β analog of a natural amino acid.

84. (New) The method of claim 8, wherein the framework is an amino acid with an amine, hydroxyl, phenol, carboxyl, thiol, thioether, or amidino group on its side chain.

85. (New) The method of claim 8, wherein the framework is a serine, threonine, tyrosine, aspartic acid, glutamic acid, asparagine, glutamine, cysteine, lysine, arginine, or histidine moiety.

86. (New) The method of claim 8, wherein the building block molecule further comprises a linker, and

the linker is a alkyl, substituted alkyl, cycloalkyl, heterocyclic, substituted heterocyclic, aryl alkyl, aryl, heteroaryl, heteroaryl alkyl, ethoxy or propoxy oligomer, or glycoside moiety; substituted with a carboxyl, alcohol, phenol, thiol, amine, carbonyl, or maleimide group.

87. (Currently Amended) The method of claim 86, wherein the linker is of the formula $(\text{CH}_2)_n\text{C}(\text{O})-$, with $n=1-16$.